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### NOTICE OF ALLOWANCE AND FEE(S) DUE

22428

7590

08/07/2009

FOLEY AND LARDNER LLP SUITE 500 3000 K STREET NW WASHINGTON, DC 20007

EXAMINER				
KOSAR, AARON J				
ART UNIT	PAPER NUMBER			

1651

DATE MAILED: 08/07/2009

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
10/591,241	08/31/2006	Ichiro Iwai	053466-0450	9322

TITLE OF INVENTION: METHOD FOR DETERMINING THE DEGREE OF PROTEIN OXIDATION IN A SKIN SAMPLE USING OXIDIZED PROTEIN IN STRATUM CORNEUM AS AN INDICATOR

APPLN. TYPE	SMALL ENTITY	ISSUE FEE DUE	PUBLICATION FEE DUE	PREV. PAID ISSUE FEE	TOTAL FEE(S) DUE	DATE DUE
nonprovisional	NO	\$1510	\$300	\$0	\$1810	11/09/2009

THE APPLICATION IDENTIFIED ABOVE HAS BEEN EXAMINED AND IS ALLOWED FOR ISSUANCE AS A PATENT. PROSECUTION ON THE MERITS IS CLOSED. THIS NOTICE OF ALLOWANCE IS NOT A GRANT OF PATENT RIGHTS. THIS APPLICATION IS SUBJECT TO WITHDRAWAL FROM ISSUE AT THE INITIATIVE OF THE OFFICE OR UPON PETITION BY THE APPLICANT. SEE 37 CFR 1.313 AND MPEP 1308.

THE ISSUE FEE AND PUBLICATION FEE (IF REQUIRED) MUST BE PAID WITHIN THREE MONTHS FROM THE MAILING DATE OF THIS NOTICE OR THIS APPLICATION SHALL BE REGARDED AS ABANDONED. THIS STATUTORY PERIOD CANNOT BE EXTENDED. SEE 35 U.S.C. 151. THE ISSUE FEE DUE INDICATED ABOVE DOES NOT REFLECT A CREDIT FOR ANY PREVIOUSLY PAID ISSUE FEE IN THIS APPLICATION. IF AN ISSUE FEE HAS PREVIOUSLY BEEN PAID IN THIS APPLICATION (AS SHOWN ABOVE), THE RETURN OF PART B OF THIS FORM WILL BE CONSIDERED A REQUEST TO REAPPLY THE PREVIOUSLY PAID ISSUE FEE TOWARD THE ISSUE FEE NOW DUE.

#### HOW TO REPLY TO THIS NOTICE:

I. Review the SMALL ENTITY status shown above.

If the SMALL ENTITY is shown as YES, verify your current SMALL ENTITY status:

A. If the status is the same, pay the TOTAL FEE(S) DUE shown above.

B. If the status above is to be removed, check box 5b on Part B - Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and twice the amount of the ISSUE FEE shown above, or

If the SMALL ENTITY is shown as NO:

A. Pay TOTAL FEE(S) DUE shown above, or

B. If applicant claimed SMALL ENTITY status before, or is now claiming SMALL ENTITY status, check box 5a on Part B - Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and 1/2 the ISSUE FEE shown above.

II. PART B - FEE(S) TRANSMITTAL, or its equivalent, must be completed and returned to the United States Patent and Trademark Office (USPTO) with your ISSUE FEE and PUBLICATION FEE (if required). If you are charging the fee(s) to your deposit account, section "4b" of Part B - Fee(s) Transmittal should be completed and an extra copy of the form should be submitted. If an equivalent of Part B is filed, a request to reapply a previously paid issue fee must be clearly made, and delays in processing may occur due to the difficulty in recognizing the paper as an equivalent of Part B.

III. All communications regarding this application must give the application number. Please direct all communications prior to issuance to Mail Stop ISSUE FEE unless advised to the contrary.

IMPORTANT REMINDER: Utility patents issuing on applications filed on or after Dec. 12, 1980 may require payment of maintenance fees. It is patentee's responsibility to ensure timely payment of maintenance fees when due.

### PART B - FEE(S) TRANSMITTAL

### Complete and send this form, together with applicable fee(s), to: Mail Mail Stop ISSUE FEE

Commissioner for Patents P.O. Box 1450 Alexandria, Virginia 22313-1450

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INSTRUCTIONS: This form should be used for transmitting the ISSUE FEE and PUBLICATION FEE (if required). Blocks 1 through 5 should be completed where ar in m

appropriate. All further indicated unless correcte maintenance fee notifica	correspondence includir ed below or directed oth	ng the Patent, advance on herwise in Block 1, by (a	rders and notification of n a) specifying a new corres	naintenance fees v pondence address:	vill be and/o	mailed to the current r (b) indicating a sepa	correspondence address as arate "FEE ADDRESS" for
		ock 1 for any change of address)	Fee(	s) Transmittal. Thers. Each additiona	is certi: I papei	ficate cannot be used f c, such as an assignme	or domestic mailings of the for any other accompanying nt or formal drawing, must
22428	7590 08/07	72009	have			iling or transmission.	
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WASHINGTON	I, DC 20007						(Depositor's name)
							(Signature)
							(Date)
APPLICATION NO.	FILING DATE		FIRST NAMED INVENTOR		ATTC	RNEY DOCKET NO.	CONFIRMATION NO.
10/591,241	08/31/2006	•	Ichiro Iwai		•	053466-0450	9322
TITLE OF INVENTION PROTEIN IN STRATUM			GREE OF PROTEIN OX	IDATION IN A	SKIN	SAMPLE USING O	XIDIZED
APPLN, TYPE	SMALL ENTITY	ISSUE FEE DUE	PUBLICATION FEE DUE	PREV. PAID ISSU	E FEE	TOTAL FEE(S) DUE	DATE DUE
nonprovisional	NO	\$1510	\$300	\$0		\$1810	11/09/2009
EXAM	IINER	ART UNIT	CLASS-SUBCLASS				
KOSAR, A		1651	435-004000				
<ol> <li>Change of correspondence address or indication of "Fee Address" (37 CFR 1.363).</li> <li>Change of correspondence address (or Change of Correspondence Address form PTO/SB/122) attached.</li> <li>"Fee Address" indication (or "Fee Address" Indication form PTO/SB/47; Rev 03-02 or more recent) attached. Use of a Customer Number is required.</li> </ol>			or agents OR, alternative (2) the name of a single registered attorney or a 2 registered patent attorney on a 2 registered patent attorney or a 3 registered patent attorney or 3 registered patent attorney or 3	of a single firm (having as a member a torney or agent) and the names of up to patent attorneys or agents. If no name is			
PLEASE NOTE: Uni recordation as set fort (A) NAME OF ASSIG	less an assignee is ident h in 37 CFR 3.11. Comp GNEE	ified below, no assignee oletion of this form is NO	T a substitute for filing an  (B) RESIDENCE: (CITY	atent. If an assign assignment. and STATE OR C	COUNT	TRY)	ocument has been filed for
Please check the appropr	iate assignee category or	categories (will not be pr	rinted on the patent):	Individual 🖵 Co	orporat	ion or other private gro	oup entity Government
4a. The following fee(s) are submitted:  ☐ Issue Fee ☐ Publication Fee (No small entity discount permitted) ☐ Advance Order - # of Copies			4b. Payment of Fee(s): (Please first reapply any previously paid issue fee shown above)  ☐ A check is enclosed. ☐ Payment by credit card. Form PTO-2038 is attached. ☐ The Director is hereby authorized to charge the required fee(s), any deficiency, or credit any overpayment, to Deposit Account Number (enclose an extra copy of this form).				
5. Change in Entity Sta	<b>tus</b> (from status indicate is SMALL ENTITY statu		b. Applicant is no long	per claiming SMA	LLEN	TITY status. See 37 CF	FR 1.27(g)(2).
NOTE: The Issue Fee an	d Publication Fee (if req		d from anyone other than t				ne assignee or other party in
Authorized Signature				Date			
This collection of inform an application. Confiden submitting the complete this form and/or suggest Box 1450, Alexandria, V Alexandria, Virginia 223	tiality is governed by 35 d application form to the ions for reducing this bu. Virginia 22313-1450. DC	CFR 1.311. The informatic U.S.C. 122 and 37 CFR USPTO. Time will vary rden, should be sent to the ONOT SEND FEES OR	on is required to obtain or r 1.14. This collection is est depending upon the indiv e Chief Information Office COMPLETED FORMS TO	etain a benefit by t imated to take 12 t idual case. Any co rr, U.S. Patent and D THIS ADDRESS	he pub minute ommen Trader S. SEN	lic which is to file (and s to complete, includin ts on the amount of tir mark Office, U.S. Depp D TO: Commissioner	by the USPTO to process) g gathering, preparing, and me you require to complete artment of Commerce, P.O. for Patents, P.O. Box 1450,

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10/591,241 08/31/2006		Ichiro Iwai	053466-0450	9322
22428 75	90 08/07/2009		EXAM	INER
FOLEY AND LARDNER LLP			KOSAR, A	AARON J
SUITE 500			ART UNIT	PAPER NUMBER
3000 K STREET NW WASHINGTON, DC 20007		1651 DATE MAILED; 08/07/200	9	

## Determination of Patent Term Adjustment under 35 U.S.C. 154 (b)

(application filed on or after May 29, 2000)

The Patent Term Adjustment to date is 306 day(s). If the issue fee is paid on the date that is three months after the mailing date of this notice and the patent issues on the Tuesday before the date that is 28 weeks (six and a half months) after the mailing date of this notice, the Patent Term Adjustment will be 306 day(s).

If a Continued Prosecution Application (CPA) was filed in the above-identified application, the filing date that determines Patent Term Adjustment is the filing date of the most recent CPA.

Applicant will be able to obtain more detailed information by accessing the Patent Application Information Retrieval (PAIR) WEB site (http://pair.uspto.gov).

Any questions regarding the Patent Term Extension or Adjustment determination should be directed to the Office of Patent Legal Administration at (571)-272-7702. Questions relating to issue and publication fee payments should be directed to the Customer Service Center of the Office of Patent Publication at 1-(888)-786-0101 or (571)-272-4200.

	Application No.	Applicant(s)
	10/591,241	IWAI ET AL.
Notice of Allowability	Examiner	Art Unit
	AARON J. KOSAR	1651
The MAILING DATE of this communication appeal all claims being allowable, PROSECUTION ON THE MERITS IS herewith (or previously mailed), a Notice of Allowance (PTOL-85) NOTICE OF ALLOWABILITY IS NOT A GRANT OF PATENT R	(OR REMAINS) CLOSED in or other appropriate commur IGHTS. This application is subsected MPEP 1308.	this application. If not included nication will be mailed in due course. <b>THIS</b>
1. This communication is responsive to the amendment filed	<u>18 March 2009</u> .	
2. ☑ The allowed claim(s) is/are <u>1 and 2</u> .		
<ul> <li>3.  Acknowledgment is made of a claim for foreign priority ur</li> <li>a)  All b)  Some* c)  None of the:</li> <li>1.  Certified copies of the priority documents have</li> <li>2.  Certified copies of the priority documents have</li> <li>3.  Copies of the certified copies of the priority documents have</li> <li>International Bureau (PCT Rule 17.2(a)).</li> <li>* Certified copies not received:</li> </ul>	been received. been received in Application	No
Applicant has THREE MONTHS FROM THE "MAILING DATE" noted below. Failure to timely comply will result in ABANDONM THIS THREE-MONTH PERIOD IS NOT EXTENDABLE.		a reply complying with the requirements
<ol> <li>A SUBSTITUTE OATH OR DECLARATION must be subm INFORMAL PATENT APPLICATION (PTO-152) which give</li> </ol>		
<ol> <li>CORRECTED DRAWINGS (as "replacement sheets") must</li> <li>(a) ☐ including changes required by the Notice of Draftspers</li> <li>1) ☐ hereto or 2) ☐ to Paper No./Mail Date</li> <li>(b) ☐ including changes required by the attached Examiner's Paper No./Mail Date</li> <li>Identifying indicia such as the application number (see 37 CFR 1 each sheet. Replacement sheet(s) should be labeled as such in t</li> <li>DEPOSIT OF and/or INFORMATION about the depo attached Examiner's comment regarding REQUIREMENT</li> </ol>	son's Patent Drawing Review  s Amendment / Comment or i  84(c)) should be written on the he header according to 37 CFF sit of BIOLOGICAL MATE	n the Office action of e drawings in the front (not the back) of t 1.121(d). RIAL must be submitted. Note the
Attachment(s)  1. ☐ Notice of References Cited (PTO-892)  2. ☐ Notice of Draftperson's Patent Drawing Review (PTO-948)  3. ☐ Information Disclosure Statements (PTO/SB/08), Paper No./Mail Date  4. ☐ Examiner's Comment Regarding Requirement for Deposit of Biological Material	6. ☐ Interview Su Paper No./N 7. ☑ Examiner's A	ormal Patent Application mmary (PTO-413), Mail Date Amendment/Comment Statement of Reasons for Allowance
	3 Suisi	

DETAILED ACTION

Claims 1-2 have been examined on the merits and found allowable (as amended within

the Examiner's Amendment below).

**EXAMINER'S AMENDMENT** 

An examiner's amendment to the record appears below. Should the changes and/or

additions be unacceptable to applicant, an amendment may be filed as provided by 37 CFR

1.312. To ensure consideration of such an amendment, it MUST be submitted no later than the

payment of the issue fee.

Authorization for this examiner's amendment was given in a telephone interview with

Stephen Maebius on July 29, 2009. Accordingly, any rejection and/or objection not specifically

addressed is hererin withdrawn.

In the Claims:

Claims 3 and 4 are canceled.

Claims 1 and 2 have been amended as follows:

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1. A method of determining the degree of protein oxidation in a skin sample comprising stratum

corneum, the method comprising:

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Art Unit: 1651

(a) providing a skin sample (S) selected from the group comprising:

- (i) stratum corneum having carbonylated protein; or
- (ii) oxidized stratum corneum, wherein the oxidized stratum corneum is obtained by contacting a skin sample comprising stratum corneum with an oxidizing agent and wherein the oxidizing agent comprises acrolein, thereby forming an acrolein-protein adduct;

wherein the skin sample (S) comprising said stratum corneum (i) or said oxidized stratum corneum (ii) is contacted and fluorescently labeled with a hydrazine fluorophore, the hydrazine fluorophore selected from the group consisting of fluorescein-thiosemicarbazide; Texas Red hydrazide; biotin hydrazide reacted with fluorescein avidin; dintrophenylhydrazine (DNPH) reacted with anti-DNP antibody and with a fluorescein-labeled secondary antibody specific to the anti-DNP antibody; and DNPH reacted with a fluorescent dye;

- (b) measuring the fluorescence of the skin sample (S); and then
- (c) determining the degree of protein oxidation of the skin sample (S),
  - (i) wherein the fluorescence intensity of the skin sample (S) is indicative of the degree of protein oxidation in the skin sample (S);
  - (ii) wherein, relative to the fluorescence intensity of a skin sample not treated with acrolein (S'), an increased fluorescence intensity of the skin sample (S) indicates an increased protein oxidation in the skin sample (S); and
  - (iii) wherein said skin sample (S) has increased protein oxidation.

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Art Unit: 1651

2. The method of claim 1, further comprising:

(d) measuring the conductance of a skin sample (S") comprising stratum corneum, wherein the skin sample (S") is oxidized by contacting with an oxidizing agent, wherein said oxidizing agent is selected from the group consisting of hypochlorous acid and acrolein, and wherein an increase in protein oxidation provides a decrease in conductance of the skin sample (S") relative to the conductance of a skin sample not contacted with an oxidizing

--

agent (S').

#### Conclusion

Any inquiry concerning this communication or earlier communications from the examiner should be directed to AARON J. KOSAR whose telephone number is (571)270-3054. The examiner can normally be reached on Monday-Thursday, 7:30AM-5:00PM, ALT. Friday, EST.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Mike Wityshyn can be reached on (571) 272-0926. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

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Art Unit: 1651

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would like assistance from a USPTO Customer Service Representative or access to the automated information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

/Aaron J Kosar/ Examiner, Art Unit 1651 /Christopher R. Tate/ Primary Examiner, Art Unit 1655